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Application No.: 10/718,461

Office Action Dated: August 13, 2004

This listing of claims will replace all prior versions, and listings, of claims in the application.

Listing of Claims:

1. (original) A compound of formulae (I) or (II) having the structure

wherein

 R_1 , R_2 , R_3 , R_4 , R_7 , R_8 , R_9 , R_{10} , R_{11} , R_{12} , R_{14} , and R_{15} are each, independently, hydrogen, R_{17} , monofluoroalkyl, monofluoroalkenyl, aryl- R_{16} -, heteroaryl- R_{16} -, hydroxyalkyl, HO- R_{16} -, R_{17} -X- R_{16} -, HS- R_{16} -, R_{17} -S(O)-, R_{17} -S(O)₂-, R_{17} -SO₃-, R_{17} -S(O)₂NR-, -N(R)₂, -NR-C(NH₂)=NR, cyano, nitro, halogen, -OR, -SR, -SO₃R, -S(O)₂N(R)₂, -C(O)R, -C(R)=N-OR, -C(NH₂)=NR, -CO₂R, -OC(O)R, or -C(O)N(R)₂; or are taken together with either R_{p+1} or R_{p-1} linked with an -alkylene-, or -X-alkylene- group;

R₅ is hydrogen, R₁₇, monofluoroalkyl, monofluoroalkenyl, aryl-R₁₆-, heteroaryl-R₁₆-, hydroxyalkyl, HO-R₁₆-, R₁₇-X-R₁₆-, HS-R₁₆-, -CR(O), -CO₂R, or -C(O)N(R)₂; or R₅ may be taken together with either R₆ or R₇ and linked with an -alkylene- or -X-alkylene- group;

 R_6 is hydrogen, R_{17} , monofluoroalkyl, monofluoroalkenyl, aryl- R_{16} -, heteroaryl- R_{16} -, hydroxyalkyl, HO- R_{16} -, R_{17} -X- R_{16} -, HS- R_{16} -, -CR(O), -CO₂R, or -C(O)N(R)₂; or R_6

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may be taken together with either R₅ or R₇ and linked with an -alkylene- or -X-alkylene- group;

- R_{13} is R, R_{17} -X- R_{16} -, R_{17} -S(O)-, R_{17} -S(O)₂-, -SO₃R, -S(O)₂N(R)₂, or D-glucuronidate;
- R₁₆ is -alkylene-, -cycloalkylene-, -alkylene-X-alkylene-, -alkylene-X-cycloalkylene-, -cycloalkylene-X-alkylene-, or -cycloalkylene-X-cycloalkylene-;
- R₁₇ is alkyl, aryl, heteroaryl, cycloalkyl, alkenyl, cycloalkenyl, alkynyl, alkenyl-X-alkylene-, cycloalkenyl-X-alkylene-, or perfluoroalkyl;
- R is, independently, hydrogen, alkyl, alkenyl, alkynyl, cycloalkyl, cycloalkenyl, monofluoroalkyl, perfluoroalkyl, aryl, arylalkyl, heteroaryl, heteroarylalkyl, hydroxy-(C₂-C₆)alkyl, alkoxyalkyl, alkylthioalkyl, formyl, acyl, alkoxycarbonyl, -C(O)NH₂, alkylaminocarbonyl, dialkylaminocarbonyl, alkylaminoalkyl, or dialkylaminoalkyl; or when an atom contains two R groups, the R groups may be taken together linked with an -alkylene- group;

X is O, -NR-, -S(O)_m-, -C(O)-, -OC(O)-, -C(O)O-, -NRC(O)-, or -C(O)NR-; m is 0, 1, or 2;

p is 2, 3, 6, 7, 8, 9, 12, 13, or 14;

- R_{21} , R_{22} , R_{23} , R_{24} , R_{27} , R_{28} , R_{29} , R_{30} , R_{31} , R_{33} , R_{34} , and R_{35} are, independently, hydrogen, R_{17} , monofluoroalkyl, monofluoroalkenyl, aryl- R_{16} -, heteroaryl- R_{16} -, hydroxyalkyl, HO- R_{16} -, R_{17} -Y- R_{16} -, HS- R_{16} -, R_{17} -S(O)-, R_{17} -S(O)₂-, R_{17} -SO₃-, R_{17} -S(O)₂NR-, -N(R)₂, -NR-C(NH₂)=NR, cyano, nitro, halogen, -OR, -SR, -SO₃R, -S(O)₂N(R)₂, -C(O)R, -C(R)=N-OR, -C(NH₂)=NR, -CO₂R, -OC(O)R, or -C(O)N(R)₂; or are taken together with either R_{q+1} or R_{q-1} linked with an -alkylene-, or -Y-alkylene- group;
- R₂₅ is hydrogen, R₁₇, monofluoroalkyl, monofluoroalkenyl, aryl-R₁₆-, heteroaryl-R₁₆-, hydroxyalkyl, HO-R₁₆-, R₁₇-Y-R₁₆-, HS-R₁₆-, -CR(O), -CO₂R, or -C(O)N(R)₂; or R₂₅ may be taken together with either R₂₆ or R₂₇ and linked with an -alkylene- or -Y-alkylene- group;
- R₂₆ is hydrogen, R₁₇, monofluoroalkyl, monofluoroalkenyl, aryl-R₁₆-, heteroaryl-R₁₆-, hydroxyalkyl, HO-R₁₆-, R₁₇-Y-R₁₆-, HS-R₁₆-, -CR(O), -CO₂R, or -C(O)N(R)₂; or R₂₆ may be taken together with either R₂₅ or R₂₇ and linked with an -alkylene- or -Y-alkylene- group;
- R_{32} is R, R_{17} -Y- R_{16} -, R_{17} -S(O)-, R_{17} -S(O)₂-, -SO₃R, -S(O)₂N(R)₂, or D-glucuronidate; Page 3 of 32

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Y is O, -NR-, -S(O)_n-, -C(O)-, -OC(O)-, -C(O)O-, -NRC(O)-, or -C(O)NR-;

n is 0, 1, or 2;

q is 22, 23, 26, 27, 28, 29, 32, 33, or 34;

or a pharmaceutically acceptable salt thereof.

- (original) The compound according to claim 1, wherein the compound is of formula
 (I) or a pharmaceutical acceptable salt thereof.
- 3. (original) The compound according to claim 2, wherein R₁₃ is hydrogen, or a pharmaceutically acceptable salt thereof.
- 4. (original) The compound according to claim 3, wherein

 $R_{1}, R_{2}, R_{3}, R_{4}, R_{7}, R_{8}, R_{9}, R_{10}, R_{11}, R_{12}, R_{14}, \text{ and } R_{15} \text{ are each, independently, hydrogen, } R_{17}, \\ aryl-R_{16}-, R_{17}-X-R_{16}-, \text{ hydroxyalkyl, HO-R}_{16}-, \text{ halogen, -OR, -COR, or -CO}_{2}R;$

 R_5 and R_6 are each, independently, hydrogen or R_{17} ;

R₁₆ is -alkylene-;

R₁₇ is alkyl, aryl, heteroaryl, or perfluoroalkyl;

R is hydrogen or alkyl; or a pharmaceutically acceptable salt thereof.

- (original) The compound according to claim 1, wherein the compound is of formula
 (II) or a pharmaceutical acceptable salt thereof.
- 6. (original) The compound according to claim 5, wherein R_{32} is hydrogen, or a pharmaceutically acceptable salt thereof.
- 7. (original) The compound according to claim 6, wherein $R_{21}, R_{22}, R_{23}, R_{24}, R_{27}, R_{28}, R_{29}, R_{30}, R_{31}, R_{33}, R_{34}, \text{ and } R_{35} \text{ are each, independently, hydrogen,} \\ R_{17}, \text{aryl-}R_{16}, R_{17}\text{-Y-}R_{16}\text{-, hydroxyalkyl, HO-}R_{16}\text{-, halogen, -OR, -COR, or -CO}_2R;$

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R₂₅ and R₂₆ are each, independently, hydrogen or R₁₇;

R₁₆ is -alkylene-;

R₁₇ is alkyl, aryl, heteroaryl, or perfluoroalkyl;

R is hydrogen or alkyl; or a pharmaceutically acceptable salt thereof.

- 8. (currently amended) The compound according to claim 1, which is
- a) 4-[(6-methylphenanthridin-5(6H)-yl)sulfonyl]phenol;
- b) $4-\{[(S)-6-methylphenanthridin-5(6H)-yl]sulfonyl\}$ phenol;
- e) $4-\{[(R)-6-\text{methylphenanthridin}-5(6H)-yl]\text{sulfonyl}\}$ phenol;
- d) 4-[(2-bromo-6-methylphenanthridin-5(6H)-yl)sulfonyl]phenol;
- e) 2-methyl-4-[(6-methylphenanthridin-5(6H)-yl)sulfonyl]phenol;
- 4-[(2-bromo-6-methylphenanthridin-5(6H)-yl)sulfonyl]-2-methylphenol;
- g) 4-[(6-butylphenanthridin-5(6H)-yl)sulfonyl]phenol;
- h) 4-[(2-bromo-6-butylphenanthridin-5(6H)-yl)sulfonyl]phenol;
- i) 4-[(6-phenylphenanthridin-5(6H)-yl)sulfonyl]phenol;
- $\frac{1}{2}$ 4-{[(S)-6-phenylphenanthridin-5(6H)-yl]sulfonyl}phenol;
- k) $4-\{[(R)-6-phenylphenanthridin-5(6H)-yl]sulfonyl\}phenol;$
- 4-[(2-bromo-6-phenylphenanthridin-5(6H)-yl)sulfonyl]phenol;
- m) 2-bromo-4-[(2-bromo-6-phenylphenanthridin-5(6H)-yl)sulfonyl]phenol;
- n) 4-[(6-tert-butylphenanthridin-5(6H)-yl)sulfonyl]phenol;
- e) $4-\{[(R)-6-tert-butylphenanthridin-5(6H)-yl]sulfonyl\}$ phenol;
- p) $4-\{[(S)-6-tert-butylphenanthridin-5(6H)-yl]sulfonyl\}$ phenol;
- 4-[(2-bromo-6-tert-butylphenanthridin-5(6H)-yl)sulfonyl]phenol;
- r) 4-[(6-ethylphenanthridin-5(6H)-yl)sulfonyl]phenol;
- 4-[(2-bromo-6-ethylphenanthridin-5(6H)-yl)sulfonyl]phenol;
- t) 4-[(6-ethylphenanthridin-5(6H)-yl)sulfonyl]-2-methylphenol;
- u) 4-[(2-bromo-6-ethylphenanthridin-5(6H)-yl)sulfonyl]-2-methylphenol;
- + 4-{[(S*)-6-[(R*)-1-methylpropyl]phenanthridin-5(6H)-yl]sulfonyl}phenol;
- w) 4-[(6-methylphenanthridin-5(6H)-yl)sulfonyl]benzene-1,2-diol;

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2-hydroxy-5-[(6-methylphenanthridin-5(6H)-yl)sulfonyl]benzoic acid;
<del>x)</del>
            ethyl 2-ethoxy-5-[(6-methylphenanthridin-5(6H)-yl)sulfonyl]benzoate;
<del>y)</del>
            2-(hydroxymethyl)-4-[(6-methylphenanthridin-5(6H)-yl)sulfonyl]phenol;
<del>z)</del>
            2-hydroxy-5-[(6-methylphenanthridin-5(6H)-yl)sulfonyl]benzaldehyde;
aa)
            4-[(6-ethyl-2-thien-3-ylphenanthridin-5(6H)-yl)sulfonyl]phenol;
bb)
            4-{[6-ethyl-2-(3-methoxyphenyl)phenanthridin-5(6H)-yl]sulfonyl}phenol;
<del>cc)</del>
            3-{6-ethyl-5-[(4-hydroxyphenyl)sulfonyl]-5,6-dihydrophenanthridin-2-yl}phenol;
dd)
            4-[(2-dibenzo[b,d]furan-4-yl-6-ethylphenanthridin-5(6H)-yl)sulfonyl]phenol;
ee)
            4-[(8-fluoro-6-methylphenanthridin-5(6H)-yl)sulfonyl]phenol;
<del>ff)</del>
             4-\{[(S)-8-fluoro-6-methylphenanthridin-5(6H)-yl] sulfonyl\} phenol; \\
gg)
            4-\{[(R)-8-fluoro-6-methylphenanthridin-5(6H)-yl]sulfonyl\}phenol;
hh)
            4-[(8-fluoro-6-methylphenanthridin-5(6H)-yl)sulfonyl]-2-methylphenol;
<del>ii)</del>
            5-[(4-hydroxyphenyl)sulfonyl]-6-methyl-5,6-dihydrophenanthridin-9-ol;
<del>jj)</del>
            5-[(4-hydroxy-3-methylphenyl)sulfonyl]-6-methyl-5,6-dihydrophenanthridin-9-ol;
kk)
             5-[(4-hydroxy-3-methylphenyl)sulfonyl]-6-methyl-5,6-dihydrophenanthridin-7-ol;
11)
             5-[(4-hydroxyphenyl)sulfonyl]-6-methyl-5,6-dihydrophenanthridin-7-ol;
 mm)
             4-[(6-ethyl-8-fluorophenanthridin-5(6H)-yl)sulfonyl]phenol;
 <del>nn)</del>
             4-[(6-ethyl-8-fluorophenanthridin-5(6H)-yl)sulfonyl]-2-methylphenol;
 <del>00)</del>
             4-[(6-ethyl-7-methylphenanthridin-5(6H)-yl)sulfonyl]phenol;
 <del>pp)</del>
             4-[(6-ethyl-9-methylphenanthridin-5(6H)-yl)sulfonyl]phenol;
 <del>99)</del>
             4-[(2-bromo-6-ethyl-8-fluorophenanthridin-5(6H)-yl)sulfonyl]phenol;
 <del>11)</del>
             4-[(2-bromo-8-fluoro-6-methylphenanthridin-5(6H)-yl)sulfonyl]phenol;
 <del>ss)</del>
             2-chloro-4-[(6-ethyl-8-fluorophenanthridin-5(6H)-yl)sulfonyl]phenol;
 <del>tt)</del>
             4-[(6-ethyl-8-fluoro-2-phenylphenanthridin-5(6H)-yl)sulfonyl]phenol;
 <del>uu)</del>
             3-[(8-fluoro-6-methylphenanthridin-5(6H)-yl)sulfonyl]phenol;
 ₩)
             2-fluoro-4-[(8-fluoro-6-methylphenanthridin-5(6H)-yl)sulfonyl]phenol;
 ₩₩)
              4-[(8-fluoro-6-methylphenanthridin-5(6H)-yl)sulfonyl]benzene-1,2-diol;
 XX)
              4-[(6-ethyl-8-fluoro-2-methylphenanthridin-5(6H)-yl)sulfonyl]phenol;
 <del>уу)</del>
              4-[(6-ethyl-8-fluoro-2-thien-3-ylphenanthridin-5(6H)-yl)sulfonyl]phenol;
  <del>22)</del>
              4-[(6-ethyl-8-fluorophenanthridin-5(6H)-yl)sulfonyl]phenyl 3,3-
  <del>aaa)</del>
              dimethylbutanoate;
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PATENT

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4-[(6-ethyl-8-fluorophenanthridin-5(6H)-yl)sulfonyl]phenyl propionate; bbb) 4-[(6-ethyl-8-fluorophenanthridin-5(6H)-yl)sulfonyl]phenyl benzoate: ccc) 2-fluoro-4-[(6-methylphenanthridin-5(6H)-yl)sulfonyl]phenol; ddd) 4-[(2-bromo-6-methylphenanthridin-5(6H)-yl)sulfonyl]-2-fluorophenol; eee) 4-[(6-methylphenanthridin-5(6H)-yl)sulfonyl]-2-(trifluoromethyl)phenol; M) 2,6-dimethyl-4-[(6-methylphenanthridin-5(6H)-yl)sulfonyl]phenol; ggg) hhh) 4-[(6,8-dimethylphenanthridin-5(6H)-yl)sulfonyl]phenol; iii) 4-[(8-chloro-6-methylphenanthridin-5(6H)-yl)sulfonyl]phenol; 4-[(2-bromo-8-chloro-6-methylphenanthridin-5(6H)-yl)sulfonyl]phenol; iii) kkk) 2-{6-ethyl-5-[(4-hydroxyphenyl)sulfonyl]-5,6-dihydrophenanthridin-2-yl}phenol; 4-{[6-ethyl-2-[4-(methylthio)phenyl]phenanthridin-5(6H)-yl]sulfonyl}phenol; HI) $4-\{[6-\text{ethyl-}2-[(E)-2-\text{phenylethenyl}]\text{phenanthridin-}5(6H)-yl]\text{sulfonyl}\}\text{phenol};$ mmm) 4-{[2-(1,1'-biphenyl-4-yl)-6-ethylphenanthridin-5(6H)-yl]sulfonyl}phenol; nnn) 4-{[2-(3-chlorophenyl)-6-ethylphenanthridin-5(6H)-yl]sulfonyl} phenol; 000) 4-[(6-ethyl-2-quinolin-8-ylphenanthridin-5(6H)-yl)sulfonyl]phenol; ppp) 4-[(6-ethyl-2-phenylphenanthridin-5(6H)-yl)sulfonyl]phenol; qqq) 4-{[6-ethyl-2-(2-methylphenyl)phenanthridin-5(6H)-yl]sulfonyl}phenol; m) 4-[(6-ethyl-2-thianthren-1-ylphenanthridin-5(6H)-yl)sulfonyl]phenol; sss) ttt) 4-{[2-(1-benzofuran-2-yl)-6-ethylphenanthridin-5(6H)-yl]sulfonyl}phenol; 4-{[6-ethyl-2-(4-hydroxyphenyl)phenanthridin-5(6H)-yl]sulfonyl}phenol; uuu) 4-{[2-(2-chlorophenyl)-6-ethylphenanthridin-5(6H)-yl]sulfonyl} phenol; ***) 4-{[6-ethyl-2-(4-ethylphenyl)phenanthridin-5(6H)-yl]sulfonyl}phenol; www) 1-(5-{6-ethyl-5-[(4-hydroxyphenyl)sulfonyl]-5,6-dihydrophenanthridin-2-XXX) yl}thien-2-yl)ethanone; 5-{6-ethyl-5-[(4-hydroxyphenyl)sulfonyl]-5,6-dihydrophenanthridin-2-ууу) yl}pyrimidine-2,4-diol; 4-{[6-ethyl-2-(2-hydroxyphenyl)phenanthridin-5(6H)-yl]sulfonyl}-2-222) methylphenol; 4-[(6-ethyl-2-thien-3-ylphenanthridin-5(6H)-yl)sulfonyl]-2-methylphenol; aaaa) 4-{[6-ethyl-2-[4-(methylthio)phenyl]phenanthridin-5(6H)-yl]sulfonyl}-2bbbb) methylphenol;

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ecce) 4-{[6-ethyl-2-[(E)-2-phenylethenyl]phenanthridin-5(6H)-yl]sulfonyl}-2-methylphenol;

dddd) 4-{6-ethyl-5-[(4-hydroxy-3-methylphenyl)sulfonyl]-5,6-dihydrophenanthridin-2-yl}benzene-1,2-diol;

eeee) 4-{[2-(1,1'-biphenyl-4-yl)-6-ethylphenanthridin-5(6H)-yl]sulfonyl}-2-methylphenol;

4-{[6-ethyl-2-(3-hydroxyphenyl)phenanthridin-5(6H)-yl]sulfonyl}-2-methylphenol;

gggg) 4-{[2-(3-chlorophenyl)-6-ethylphenanthridin-5(6H)-yl]sulfonyl}-2-methylphenol;

hhhh) 4-{[6-ethyl-2-[(E)-hept-1-enyl]phenanthridin-5(6H)-yl]sulfonyl}-2-methylphenol;

4-[(6-ethyl-2-pyridin-4-ylphenanthridin-5(6H)-yl)sulfonyl]-2-methylphenol;

4-[(6-ethyl-2-quinolin-8-ylphenanthridin-5(6H)-yl)sulfonyl]-2-methylphenol;

kkkk) 4-{[6-ethyl-2-(2-methylphenyl)phenanthridin-5(6H)-yl]sulfonyl}-2-methylphenol;

4-{[2-(1-benzothien-2-yl)-6-ethylphenanthridin-5(6H)-yl]sulfonyl}-2-methylphenol;

mmmm) 4-{[2-(1-benzothien-3-yl)-6-ethylphenanthridin-5(6H)-yl]sulfonyl}-2-methylphenol;

 $\frac{1}{1}$ 4-[(2-dibenzo[b,d]furan-4-yl-6-ethylphenanthridin-5(6H)-yl)sulfonyl]-2-methylphenol;

••••• 4-{[2-(1-benzofuran-2-yl)-6-ethylphenanthridin-5(6H)-yl]sulfonyl}-2-methylphenol;

pppp) 4-{[6-ethyl-2-(4-hydroxyphenyl)phenanthridin-5(6H)-yl]sulfonyl}-2-methylphenol;

4-{[2-(2-chlorophenyl)-6-ethylphenanthridin-5(6H)-yl]sulfornyl}-2-methylphenol;

4-{[6-ethyl-2-(4-ethylphenyl)phenanthridin-5(6H)-yl]sulfonyl}-2-methylphenol;

1-(5-{6-ethyl-5-[(4-hydroxy-3-methylphenyl)sulfonyl]-5,6-dihydrophenanthridin-2-yl}thien-2-yl)ethanone;

5-{6-ethyl-5-[(4-hydroxy-3-methylphenyl)sulfonyl]-5,6-dihydrophenanthridin-2-yl}pyrimidine-2,4-diol;

4-{[(6R)-3,8-difluoro-6-methylphenanthridin-5(6H)-yl]sulfonyl}phenol;

4-{[(6S)-3,8-difluoro-6-methylphenanthridin-5(6H)-yl]sulfonyl}phenol;

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3-{[(6R)-3.8-difluoro-6-methylphenanthridin-5(6H)-yl]sulfonyl}phenol;

3-{[(6S)-3,8-difluoro-6-methylphenanthridin-5(6H)-yl]sulfonyl}phenol;

4-{[(6S)-3,8-difluoro-6-methylphenanthridin-5(6H)-yl]sulfonyl}benzene-1,3-diol;

4-{[(6R)-3,8-difluoro-6-methylphenanthridin-5(6H)-yl]sulfonyl}benzene-1,3-diol;

4-[(3-fluoro-6-methylphenanthridin-5(6H)-yl)sulfonyl]phenol;

3-[(3-fluoro-6-methylphenanthridin-5(6H)-yl)sulfonyl]phenol;

3-{[(6R)-3-fluoro-6-methylphenanthridin-5(6H)-yl]sulfonyl}phenol;

3-{[(6S)-3-fluoro-6-methylphenanthridin-5(6H)-yl]sulfonyl}phenol;

4-[3-fluoro-6-methylphenanthridin-5(6H)-yl]sulfonyl]benzene-1,3-diol;

4-{[(6R)-3-fluoro-6-methylphenanthridin-5(6H)-yl]sulfonyl}benzene-1,3-diol;

4-{[(6S)-3-fluoro-6-methylphenanthridin-5(6H)-yl]sulfonyl}benzene-1,3-diol;

4-[(2-fluoro-6-methylphenanthridin-5(6H)-yl)sulfonyl]phenol;

3-[(2-fluoro-6-methylphenanthridin-5(6H)-yl)sulfonyl]phenol;

4-[(3,9-difluoro-6-methylphenanthridin-5(6H)-yl)sulfonyl]phenol;

3-[(3,9-difluoro-6-methylphenanthridin-5(6H)-yl)sulfonyl]phenol;

4-[(2,9-difluoro-6-methylphenanthridin-5(6H)-yl)sulfonyl]phenol;

3-[(2,9-difluoro-6-methylphenanthridin-5(6H)-yl)sulfonyl]phenol;

3-[(3-chloro-6-methylphenanthridin-5(6H)-yl)sulfonyl]phenol;

3-{[(6R)-3-chloro-6-methylphenanthridin-5(6H)-yl]sulfonyl}phenol;

3-{[(6S)-3-chloro-6-methylphenanthridin-5(6H)-yl]sulfonyl}phenol;

4-[(3-chloro-6-methylphenanthridin-5(6H)-yl)sulfonyl]phenol;

4-{[(6R)-3-chloro-6-methylphenanthridin-5(6H)-yl]sulfonyl}phenol;

4-{[(6S)-3-chloro-6-methylphenanthridin-5(6H)-yl]sulfonyl}phenol;

4-{[(6R)-3-chloro-6-methylphenanthridin-5(6H)-yl]sulfonyl}benzene-1,3-diol;

4-{[(6S)-3-chloro-6-methylphenanthridin-5(6H)-yl]sulfonyl}benzene-1,3-diol;

4-{[(6S)-8-fluoro-6-methylphenanthridin-5(6H)-yl]sulfonyl}phenyl sulfamate;

4-[(6-ethyl-8-fluoro-2-pyridin-3-ylphenanthridin-5(6H)-yl)sulfonyl]phenol;

or a pharmaceutically acceptable salt thereof.

9. (original) A pharmaceutical composition, which comprises a compound of formulae (I) or (II) having the structure

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wherein

- R_1 , R_2 , R_3 , R_4 , R_7 , R_8 , R_9 , R_{10} , R_{11} , R_{12} , R_{14} , and R_{15} are each, independently, hydrogen, R_{17} , monofluoroalkyl, monofluoroalkenyl, aryl- R_{16} -, heteroaryl- R_{16} -, hydroxyalkyl, HO- R_{16} -, R_{17} -X- R_{16} -, HS- R_{16} -, R_{17} -S(O)-, R_{17} -S(O)₂-, R_{17} -SO₃-, R_{17} -S(O)₂NR-, -N(R)₂, -NR-C(NH₂)=NR, cyano, nitro, halogen, -OR, -SR, -SO₃R, -S(O)₂N(R)₂, -C(O)R, -C(R)=N-OR, -C(NH₂)=NR, -CO₂R, -OC(O)R, or -C(O)N(R)₂; or are taken together with either R_{p+1} or R_{p-1} linked with an -alkylene-, or -X-alkylene- group;
- R₅ is hydrogen, R₁₇, monofluoroalkyl, monofluoroalkenyl, aryl-R₁₆-, heteroaryl-R₁₆-, hydroxyalkyl, HO-R₁₆-, R₁₇-X-R₁₆-, HS-R₁₆-, -CR(O), -CO₂R, or -C(O)N(R)₂; or R₅ may be taken together with either R₆ or R₇ and linked with an -alkylene- or -X-alkylene- group;
- R₆ is hydrogen, R₁₇, monofluoroalkyl, monofluoroalkenyl, aryl-R₁₆-, heteroaryl-R₁₆-, hydroxyalkyl, HO-R₁₆-, R₁₇-X-R₁₆-, HS-R₁₆-, -CR(O), -CO₂R, or -C(O)N(R)₂; or R₆ may be taken together with either R₅ or R₇ and linked with an -alkylene- or -X-alkylene- group;
- R_{13} is R, R_{17} -X- R_{16} -, R_{17} -S(O)-, R_{17} -S(O)₂-, -SO₃R, -S(O)₂N(R)₂, or D-glucuronidate;
- R₁₆ is -alkylene-, -cycloalkylene-, -alkylene-X-alkylene-, -alkylene-X-cycloalkylene-, -cycloalkylene-X-alkylene-, or -cycloalkylene-X-cycloalkylene-;

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R₁₇ is alkyl, aryl, heteroaryl, cycloalkyl, alkenyl, cycloalkenyl, alkynyl, alkenyl-X-alkylene-, cycloalkenyl-X-alkylene-, or perfluoroalkyl;

R is, independently, hydrogen, alkyl, alkenyl, alkynyl, cycloalkyl, cycloalkenyl, monofluoroalkyl, perfluoroalkyl, aryl, arylalkyl, heteroaryl, heteroarylalkyl, hydroxy-(C2-C6)alkyl, alkoxyalkyl, alkylthioalkyl, formyl, acyl, alkoxycarbonyl, -C(O)NH2, alkylaminocarbonyl, dialkylaminocarbonyl, alkylaminoalkyl, or dialkylaminoalkyl, or when an atom contains two R groups, the R groups may be taken together linked with an -alkylene- group;

X is O, -NR-, -S(O)_m-, -C(O)-, -OC(O)-, -C(O)O-, -NRC(O)-, or -C(O)NR-; m is 0, 1, or 2;

p is 2, 3, 6, 7, 8, 9, 12, 13, or 14;

 R_{21} , R_{22} , R_{23} , R_{24} , R_{27} , R_{28} , R_{29} , R_{30} , R_{31} , R_{33} , R_{34} , and R_{35} are, independently, hydrogen, R_{17} ; monofluoroalkyl, monofluoroalkenyl, aryl- R_{16} -, heteroaryl- R_{16} -, hydroxyalkyl, HO- R_{16} -, R_{17} -Y- R_{16} -, HS- R_{16} -, R_{17} -S(O)-, R_{17} -S(O)₂-, R_{17} -SO₃-, R_{17} -S(O)₂NR-, -N(R)₂, -NR-C(NH₂)=NR, cyano, nitro, halogen, -OR, -SR, -SO₃R, -S(O)₂N(R)₂, -C(O)R, -C(R)=N-OR, -C(NH₂)=NR, -CO₂R, -OC(O)R, or -C(O)N(R)₂; or are taken together with either R_{q+1} or R_{q-1} linked with an -alkylene-, or -Y-alkylene- group;

R₂₅ is hydrogen, R₁₇, monofluoroalkyl, monofluoroalkenyl, aryl-R₁₆-, heteroaryl-R₁₆-, hydroxyalkyl, HO-R₁₆-, R₁₇-Y-R₁₆-, HS-R₁₆-, -CR(O), -CO₂R, or -C(O)N(R)₂; or R₂₅ may be taken together with either R₂₆ or R₂₇ and linked with an -alkylene- or -Y-alkylene- group;

R₂₆ is hydrogen, R₁₇, monofluoroalkyl, monofluoroalkenyl, aryl-R₁₆-, heteroaryl-R₁₆-, hydroxyalkyl, HO-R₁₆-, R₁₇-Y-R₁₆-, HS-R₁₆-, -CR(O), -CO₂R, or -C(O)N(R)₂; or R₂₆ may be taken together with either R₂₅ or R₂₇ and linked with an -alkylene- or -Y-alkylene- group;

 R_{32} is R, R_{17} -Y- R_{16} -, R_{17} -S(O)-, R_{17} -S(O)₂-, -SO₃R, -S(O)₂N(R)₂, or D-glucuronidate; Y is O, -NR-, -S(O)_n-, -C(O)-, -OC(O)-, -C(O)O-, -NRC(O)-, or -C(O)NR-; n is 0, 1, or 2;

q is 22, 23, 26, 27, 28, 29, 32, 33, or 34;

or a pharmaceutically acceptable salt thereof, and a pharmaceutically acceptable carrier.

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10. (currently amended) A method of treating or inhibiting chronic inflammatory disease in a mammal in need thereof, which comprises administering to said mammal an effective amount of a compound of claim 1.

- 11. (currently amended) A method of treating or inhibiting rheumatoid arthritis, spondyloarthropathies, osteoarthritis, psoriatic arthritis, or juvenile arthritis in a mammal in need thereof, which comprises administering to said mammal an effective amount of a compound of claim 1.
- 12. (currently amended) A method of treating or inhibiting inflammatory bowel disease, Crohn's disease, ulcerative colitis, or indeterminate colitis in a mammal in need thereof, which comprises administering to said mammal an effective amount of a compound of claim 1.
- 13. (currently amended) A method of treating or inhibiting psoriasis in a mammal in need thereof, which comprises administering to said mammal an effective amount of a compound of claim 1.
- 14. (currently amended) A method of treating or inhibiting asthma or chronic obstructive pulmonary disease in a mammal in need thereof, which comprises administering to said mammal an effective amount of a compound of claim 1.
- 15. (currently amended) A method of treating or inhibiting stroke, ischemia, or reperfusion injury in a mammal in need thereof, which comprises administering to said mammal an effective amount of a compound of claim 1.
- 16. (currently amended) A method of lowering cholesterol, triglycerides, Lp(a), and LDL levels; inhibiting or treating hypercholesteremia, hyperlipidemia, cardiovascular disease, atherosclerosis, acute coronary syndrome, peripheral vascular disease, restenosis, or vasospasm in a mammal in need thereof, which comprises administering to said mammal an effective amount of a compound of claim 1.

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- 17. (currently amended) A method of treating or inhibiting Alzheimer's disease, cognitive decline, or senile dementia in a mammal in need thereof, which comprises administering to said mammal an effective amount of a compound of claim 1.
- 18. (currently amended) A method of treating or inhibiting type II diabetes in a mammal in need thereof, which comprises administering to said mammal an effective amount of a compound of claim 1.
- 19. (currently amended) A method of treating or inhibiting sepsis in a mammal in need thereof, which comprises administering to said mammal an effective amount of a compound of claim 1.
- 20. (new) The compound according to claim 2, wherein R_{13} is $-S(O)_2NH_2$, or a pharmaceutically acceptable salt thereof.
- 21. (new) The compound according to claim 5, wherein R_{32} is $-S(O)_2NH_2$, or a pharmaceutically acceptable salt thereof.
- 22. (new) A process comprising providing a sulfonamide of formula 37:

wherein

R₁, R₂, R₃, R₄, R₇, R₈, R₉, R₁₀, R₁₁, R₁₂, R₁₄, and R₁₅ are each, independently, hydrogen, R₁₇, monofluoroalkyl, monofluoroalkenyl, aryl-R₁₆-, heteroaryl-R₁₆-, hydroxyalkyl, Page 13 of 32

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HO-R₁₆-, R₁₇-X-R₁₆-, HS-R₁₆-, R₁₇-S(O)-, R₁₇-S(O)₂-, R₁₇-SO₃-, R₁₇-S(O)₂NR-, -N(R)₂, -NR-C(NH₂)=NR, cyano, nitro, halogen, -OR, -SR, -SO₃R, -S(O)₂N(R)₂, -C(O)R, -C(R)=N-OR, -C(NH₂)=NR, -CO₂R, -OC(O)R, or -C(O)N(R)₂; or are taken together with either R_{p+1} or R_{p-1} linked with an -alkylene-, or -X-alkylene- group;

- R₅ is hydrogen, R₁₇, monofluoroalkyl, monofluoroalkenyl, aryl-R₁₆-, heteroaryl-R₁₆-, hydroxyalkyl, HO-R₁₆-, R₁₇-X-R₁₆-, HS-R₁₆-, -CR(O), -CO₂R, or -C(O)N(R)₂; or R₅ may be taken together with either R₆ or R₇ and linked with an -alkylene- or -X-alkylene- group;
- R₆ is hydrogen, R₁₇, monofluoroalkyl, monofluoroalkenyl, aryl-R₁₆-, heteroaryl-R₁₆-, hydroxyalkyl, HO-R₁₆-, R₁₇-X-R₁₆-, HS-R₁₆-, -CR(O), -CO₂R, or -C(O)N(R)₂; or R₆ may be taken together with either R₅ or R₇ and linked with an -alkylene- or -X-alkylene- group;
- R_{13} is R, R_{17} -X- R_{16} -, R_{17} -S(O)-, R_{17} -S(O)₂-, -SO₃R, -S(O)₂N(R)₂, or D-glucuronidate;
- R₁₆ is -alkylene-, -cycloalkylene-, -alkylene-X-alkylene-, -alkylene-X-cycloalkylene-, -cycloalkylene-X-cycloalkylene-;
- R₁₇ is alkyl, aryl, heteroaryl, cycloalkyl, alkenyl, cycloalkenyl, alkynyl, alkenyl-X-alkylene-, cycloalkenyl-X-alkylene-, or perfluoroalkyl;
- R is, independently, hydrogen, alkyl, alkenyl, alkynyl, cycloalkyl, cycloalkenyl, monofluoroalkyl, perfluoroalkyl, aryl, arylalkyl, heteroaryl, heteroarylalkyl, hydroxy-(C2-C6)alkyl, alkoxyalkyl, alkylthioalkyl, formyl, acyl, alkoxycarbonyl, -C(O)NH2, alkylaminocarbonyl, dialkylaminocarbonyl, alkylaminoalkyl, or dialkylaminoalkyl; or when an atom contains two R groups, the R groups may be taken together linked with an -alkylene- group;

X is O, -NR-, -S(O)_m-, -C(O)-, -OC(O)-, -C(O)O-, -NRC(O)-, or -C(O)NR-;

m is 0, 1, or 2; and

p is 2, 3, 6, 7, 8, 9, 12, 13, or 14; and

treating the sulfonamide of formula 37 with potassium carbonate to produce a phenanthridine of formula I:

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(new) The process of claim 22 further comprising providing the S enantiomer of the 23. biphenylamine of formula 36:

$$R_{10}$$
 R_{10}
 R

wherein

L is fluorine or chlorine; and

reacting the S enantiomer of the biphenylamine of formula 36 with a compound of formula 3 or an anhydride:

to produce a sulfonamide of formula 37.

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24. (new) The process of claim 23 further comprising providing a biphenylamine of formula 36:

$$R_{10}$$
 R_{10}
 R

separating the biphenylamine of formula 36 into its respective enantiomers.

25. (new) The process of claim 24 further comprising providing a compound of formula 35:

reacting the compound of formula 35 with an ammonium source optionally in the presence of an acid catalyst to produce an intermediate imine; and

reducing the intermediate imine with a hydride source to produce a biphenylamine of formula 36.

26. (new) The process of claim 25 further comprising providing a compound of formula33:

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(33)

wherein

R₃₆ and R₃₇ are, independently, hydrogen or (C₁-C₄) lower straight chain or (C₃-C₆) branched chain alkyl, or R₃₆ and R₃₇ are taken together to form a pinacol moiety; and reacting the compound of formula 33 in the presence of a coupling catalyst with a compound of formula 34:

wherein

W is a chlorine, bromine, or iodine atom, or a triflate (-OSO₂CF₃) moiety; to produce a compound of formula 35.

27. (new) A process for preparing a compound of formula I:

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wherein

- R_1 , R_2 , R_3 , R_4 , R_7 , R_8 , R_9 , R_{10} , R_{11} , R_{12} , R_{14} , and R_{15} are each, independently, hydrogen, R_{17} , monofluoroalkyl, monofluoroalkenyl, aryl- R_{16} -, heteroaryl- R_{16} -, hydroxyalkyl, HO- R_{16} -, R_{17} -X- R_{16} -, HS- R_{16} -, R_{17} -S(O)-, R_{17} -S(O)₂-, R_{17} -SO₃-, R_{17} -S(O)₂NR-, -N(R)₂, -NR-C(NH₂)=NR, cyano, nitro, halogen, -OR, -SR, -SO₃R, -S(O)₂N(R)₂, -C(O)R, -C(R)=N-OR, -C(NH₂)=NR, -CO₂R, -OC(O)R, or -C(O)N(R)₂; or are taken together with either R_{p+1} or R_{p-1} linked with an -alkylene-, or -X-alkylene- group;
- R₅ is hydrogen, R₁₇, monofluoroalkyl, monofluoroalkenyl, aryl-R₁₆-, heteroaryl-R₁₆-, hydroxyalkyl, HO-R₁₆-, R₁₇-X-R₁₆-, HS-R₁₆-, -CR(O), -CO₂R, or -C(O)N(R)₂; or R₅ may be taken together with either R₆ or R₇ and linked with an -alkylene- or -X-alkylene- group;
- R₆ is hydrogen, R₁₇, monofluoroalkyl, monofluoroalkenyl, aryl-R₁₆-, heteroaryl-R₁₆-, hydroxyalkyl, HO-R₁₆-, R₁₇-X-R₁₆-, HS-R₁₆-, -CR(O), -CO₂R, or -C(O)N(R)₂; or R₆ may be taken together with either R₅ or R₇ and linked with an -alkylene- or -X-alkylene- group;
- R_{13} is R, R_{17} -X- R_{16} -, R_{17} -S(O)-, R_{17} -S(O)₂-, -SO₃R, -S(O)₂N(R)₂, or D-glucuronidate;
- R₁₆ is -alkylene-, -cycloalkylene-, -alkylene-X-alkylene-, -alkylene-X-cycloalkylene-, -cycloalkylene-X-cycloalkylene-;

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R₁₇ is alkyl, aryl, heteroaryl, cycloalkyl, alkenyl, cycloalkenyl, alkynyl, alkenyl-X-alkylene-, cycloalkenyl-X-alkylene-, or perfluoroalkyl;

R is, independently, hydrogen, alkyl, alkenyl, alkynyl, cycloalkyl, cycloalkenyl, monofluoroalkyl, perfluoroalkyl, aryl, arylalkyl, heteroaryl, heteroarylalkyl, hydroxy-(C₂-C₆)alkyl, alkoxyalkyl, alkylthioalkyl, formyl, acyl, alkoxycarbonyl, -C(O)NH₂, alkylaminocarbonyl, dialkylaminocarbonyl, alkylaminoalkyl, or dialkylaminoalkyl; or when an atom contains two R groups, the R groups may be taken together linked with an -alkylene- group;

X is O, -NR-, -S(O)_m-, -C(O)-, -OC(O)-, -C(O)O-, -NRC(O)-, or -C(O)NR-; m is 0, 1, or 2; and p is 2, 3, 6, 7, 8, 9, 12, 13, or 14;

comprising

reacting a compound of formula 33:

(33)

wherein

L is fluorine or chlorine; and

R₃₆ and R₃₇ are, independently, hydrogen or (C₁-C₄) lower straight chain or (C₃-C₆) branched chain alkyl, or R₃₆ and R₃₇ are taken together to form a pinacol moiety;

in the presence of a coupling catalyst with a compound of formula 34:

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wherein

W is a chlorine, bromine, or iodine atom, or a triflate (-OSO₂CF₃) moiety;

to produce a compound of formula 35:

- reacting the compound of formula 35 with an ammonium source optionally in the presence of an acid catalyst to produce an intermediate imine;
- reducing the intermediate imine with a hydride source to produce a biphenylamine of formula 36:

$$R_{10}$$
 R_{10}
 R

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separating the biphenylamine of formula 36 into its respective enantiomers;

• reacting the S enantiomer of the biphenylamine of formula 36 with a compound of formula 3 or an anhydride:

to produce a sulfonamide of formula 37:

• treating the sulfonamide of formula 37 with potassium carbonate to produce a phenanthridine of formula I:

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28. (new) A process comprising providing a sulfonamide of formula 37a:

wherein

 R_{21} , R_{22} , R_{23} , R_{24} , R_{27} , R_{28} , R_{29} , R_{30} , R_{31} , R_{33} , R_{34} , and R_{35} are, independently, hydrogen, R_{17} , monofluoroalkyl, monofluoroalkenyl, aryl- R_{16} -, heteroaryl- R_{16} -, hydroxyalkyl, HO- R_{16} -, R_{17} -Y- R_{16} -, HS- R_{16} -, R_{17} -S(O)-, R_{17} -S(O)₂-, R_{17} -SO₃-, R_{17} -S(O)₂NR-, -N(R)₂, -NR-C(NH₂)=NR, cyano, nitro, halogen, -OR, -SR, -SO₃R, -S(O)₂N(R)₂, -C(O)R, -C(R)=N-OR, -C(NH₂)=NR, -CO₂R, -OC(O)R, or -C(O)N(R)₂; or are taken together with either R_{q+1} or R_{q+1} linked with an -alkylene-, or -Y-alkylene- group;

R₂₅ is hydrogen, R₁₇, monofluoroalkyl, monofluoroalkenyl, aryl-R₁₆-, heteroaryl-R₁₆-, hydroxyalkyl, HO-R₁₆-, R₁₇-Y-R₁₆-, HS-R₁₆-, -CR(O), -CO₂R, or -C(O)N(R)₂; or R₂₅

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may be taken together with either R_{26} or R_{27} and linked with an -alkylene- or -Y-alkylene- group;

R₂₆ is hydrogen, R₁₇, monofluoroalkyl, monofluoroalkenyl, aryl-R₁₆-, heteroaryl-R₁₆-, hydroxyalkyl, HO-R₁₆-, R₁₇-Y-R₁₆-, HS-R₁₆-, -CR(O), -CO₂R, or -C(O)N(R)₂; or R₂₆ may be taken together with either R₂₅ or R₂₇ and linked with an -alkylene- or -Y-alkylene- group;

 R_{32} is R, R_{17} -Y- R_{16} -, R_{17} -S(O)-, R_{17} -S(O)₂-, -SO₃R, -S(O)₂N(R)₂, or D-glucuronidate;

R₁₆ is -alkylene-, -cycloalkylene-, -alkylene-X-alkylene-, -alkylene-X-cycloalkylene-, -cycloalkylene-X-cycloalkylene-;

R₁₇ is alkyl, aryl, heteroaryl, cycloalkyl, alkenyl, cycloalkenyl, alkynyl, alkenyl-X-alkylene-, cycloalkenyl-X-alkylene-, or perfluoroalkyl;

R is, independently, hydrogen, alkyl, alkenyl, alkynyl, cycloalkyl, cycloalkenyl, monofluoroalkyl, perfluoroalkyl, aryl, arylalkyl, heteroaryl, heteroarylalkyl, hydroxy-(C₂-C₆)alkyl, alkoxyalkyl, alkylthioalkyl, formyl, acyl, alkoxycarbonyl, -C(O)NH₂, alkylaminocarbonyl, dialkylaminocarbonyl, alkylaminoalkyl, or dialkylaminoalkyl; or when an atom contains two R groups, the R groups may be taken together linked with an -alkylene- group;

Y is O, -NR-, -S(O)_n-, -C(O)-, -OC(O)-, -C(O)O-, -NRC(O)-, or -C(O)NR-;

n is 0, 1, or 2; and

q is 22, 23, 26, 27, 28, 29, 32, 33, or 34; and

treating the sulfonamide of formula 37a with potassium carbonate to produce a phenanthridine of formula II:

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$$R_{30}$$
 R_{29}
 R_{28}
 R_{21}
 R_{21}
 R_{25}
 R_{24}
 R_{25}
 R_{25}
 R_{31}
 R_{34}
 R_{33}
 R_{33}
(III)

29. (new) The process of claim 28 further comprising providing the S enantiomer of the biphenylamine of formula 36a:

wherein

L is fluorine or chlorine; and

reacting the S enantiomer of the biphenylamine of formula 36a with a compound of formula 32 or an anhydride:

to produce a sulfonamide of formula 37a.

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30. (new) The process of claim 29 further comprising providing a biphenylamine of formula 36a:

$$R_{20}$$
 R_{28} R_{21} R_{20} R_{20} R_{20} R_{20} R_{20} R_{20} R_{20} R_{20} R_{20} ; and

separating the biphenylamine of formula 36a into its respective enantiomers.

31. (new) The process of claim 30 further comprising providing a compound of formula 35a:

reacting the compound of formula 35a with an ammonium source optionally in the presence of an acid catalyst to produce an intermediate imine; and

reducing the intermediate imine with a hydride source to produce a biphenylamine of formula 36.

32. (new) The process of claim 31 further comprising providing a compound of formula 33a:

(33a)

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wherein

R₃₆ and R₃₇ are, independently, hydrogen or (C₁-C₄) lower straight chain or (C₃-C₆) branched chain alkyl, or R₃₆ and R₃₇ are taken together to form a pinacol moiety; and reacting the compound of formula 33a in the presence of a coupling catalyst with a compound of formula 34a:

wherein

W is a chlorine, bromine, or iodine atom, or a triflate (-OSO₂CF₃) moiety; to produce a compound of formula 35.

33. (new) A process for preparing a compound of formula II:

$$R_{20}$$
 R_{21}
 R_{21}
 R_{21}
 R_{22}
 R_{23}
 R_{24}
 R_{25}
 R_{25}
 R_{35}
 R_{34}
 R_{33}
 R_{34}
 R_{33}
(II)

wherein

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- R_{21} , R_{22} , R_{23} , R_{24} , R_{27} , R_{28} , R_{29} , R_{30} , R_{31} , R_{33} , R_{34} , and R_{35} are, independently, hydrogen, R_{17} , monofluoroalkyl, monofluoroalkenyl, aryl- R_{16} -, heteroaryl- R_{16} -, hydroxyalkyl, HO- R_{16} -, R_{17} -Y- R_{16} -, HS- R_{16} -, R_{17} -S(O)-, R_{17} -S(O)₂-, R_{17} -SO₃-, R_{17} -S(O)₂NR-, -N(R)₂, -NR-C(NH₂)=NR, cyano, nitro, halogen, -OR, -SR, -SO₃R, -S(O)₂N(R)₂, -C(O)R, -C(R)=N-OR, -C(NH₂)=NR, -CO₂R, -OC(O)R, or -C(O)N(R)₂; or are taken together with either R_{q+1} or R_{q-1} linked with an -alkylene-, or -Y-alkylene- group;
- R₂₅ is hydrogen, R₁₇, monofluoroalkyl, monofluoroalkenyl, aryl-R₁₆-, heteroaryl-R₁₆-, hydroxyalkyl, HO-R₁₆-, R₁₇-Y-R₁₆-, HS-R₁₆-, -CR(O), -CO₂R, or -C(O)N(R)₂; or R₂₅ may be taken together with either R₂₆ or R₂₇ and linked with an -alkylene- or -Y-alkylene- group;
- R₂₆ is hydrogen, R₁₇, monofluoroalkyl, monofluoroalkenyl, aryl-R₁₆-, heteroaryl-R₁₆-, hydroxyalkyl, HO-R₁₆-, R₁₇-Y-R₁₆-, HS-R₁₆-, -CR(O), -CO₂R, or -C(O)N(R)₂; or R₂₆ may be taken together with either R₂₅ or R₂₇ and linked with an -alkylene- or -Y-alkylene- group;
- R_{32} is R, R_{17} -Y- R_{16} -, R_{17} -S(O)-, R_{17} -S(O)₂-, -SO₃R, -S(O)₂N(R)₂, or D-glucuronidate;
- R₁₆ is -alkylene-, -cycloalkylene-, -alkylene-X-alkylene-, -alkylene-X-cycloalkylene-, -cycloalkylene-X-cycloalkylene-;
- R₁₇ is alkyl, aryl, heteroaryl, cycloalkyl, alkenyl, cycloalkenyl, alkynyl, alkenyl-X-alkylene-, cycloalkenyl-X-alkylene-, or perfluoroalkyl;
- R is, independently, hydrogen, alkyl, alkenyl, alkynyl, cycloalkyl, cycloalkenyl, monofluoroalkyl, perfluoroalkyl, aryl, arylalkyl, heteroaryl, heteroarylalkyl, hydroxy-(C₂-C₆)alkyl, alkoxyalkyl, alkylthioalkyl, formyl, acyl, alkoxycarbonyl, -C(O)NH₂, alkylaminocarbonyl, dialkylaminocarbonyl, alkylaminoalkyl, or dialkylaminoalkyl; or when an atom contains two R groups, the R groups may be taken together linked with an -alkylene- group;

Y is O, -NR-, -S(O)_n-, -C(O)-, -OC(O)-, -C(O)O-, -NRC(O)-, or -C(O)NR-; n is 0, 1, or 2; q is 22, 23, 26, 27, 28, 29, 32, 33, or 34;

comprising

reacting a compound of formula 33a:

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(33a)

wherein

L is fluorine or chlorine; and

R₃₆ and R₃₇ are, independently, hydrogen or (C₁-C₄) lower straight chain or (C₃-C₆) branched chain alkyl, or R₃₆ and R₃₇ are taken together to form a pinacol moiety;

in the presence of a coupling catalyst with a compound of formula 34a:

wherein

W is a chlorine, bromine, or iodine atom, or a triflate (-OSO₂CF₃) moiety;

to produce a compound of formula 35a:

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$$R_{20}$$
 R_{20}
 R_{20}
 R_{20}
 R_{20}
 R_{20}
 R_{20}
 R_{20}
 R_{20}
 R_{20}

- reacting the compound of formula 35a with an ammonium source optionally in the presence of an acid catalyst to produce an intermediate imine;
- reducing the intermediate imine with a hydride source to produce a biphenylamine of formula 36a:

- separating the biphenyl amine of formula 36a into its respective enantiomers;
- reacting the S enantiomer of the biphenylamine of formula 36a with a compound of formula 32 or an anhydride:

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to produce a sulfonamide of formula 37a:

$$R_{20}$$
 R_{21}
 R_{26}
 R_{26}
 R_{25}
 R_{24}
 R_{35}
 R_{34}
 R_{33}
 R_{33}
 R_{34}
 R_{33}
 R_{34}
 R_{33}
 R_{34}
 R_{33}
 R_{34}

• treating the sulfonamide of formula 37a with potassium carbonate to produce a phenanthridine of formula II: